=> fil reg FILE 'REGISTRY' ENTERED AT 11:11:23 ON 06 JUN 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 5 JUN 2006 HIGHEST RN 886840-90-0 DICTIONARY FILE UPDATES: 5 JUN 2006 HIGHEST RN 886840-90-0

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Structure search iteration limits have been increased. See  ${\tt HELP\ SLIMITS}$  for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

C18 H37 N5 O9 . 5/2 H2 O4 S

=> d ide can tot

MF

```
L96 ANSWER 1 OF 4 REGISTRY COPYRIGHT 2006 ACS on STN
     49842-07-1 REGISTRY
     Entered STN: 16 Nov 1984
     D-Streptamine, O-3-amino-3-deoxy-\alpha-D-glucopyranosyl-(1\rightarrow 6)-O-
     [2,6-diamino-2,3,6-trideoxy-\alpha-D-ribo-hexopyranosyl-(1\rightarrow4)]-2-
     deoxy-, sulfate (2:5) (salt) (9CI) (CA INDEX NAME)
OTHER NAMES:
     1-Epitobramycin sulfate
CN
     Distobram
CN
CN
     Gernebein
CN
     Nebcin
CN
     Nebicina
CN
     Obracin
CN
     Tenemicin
CN
     Tobi
CN
     Tobra
CN
     Tobramycin sulfate
FS
     STEREOSEARCH
DR
     79645-27-5
```

CI COM

LC STN Files: ADISNEWS, ANABSTR, BEILSTEIN\*, BIOSIS, BIOTECHNO, CA, CABA, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMLIST, CIN, CSCHEM, EMBASE, IMSPATENTS, IPA, MEDLINE, MRCK\*, MSDS-OHS, PROMT, PS, RTECS\*, SCISEARCH, TOXCENTER, USAN, USPAT2, USPATFULL

(\*File contains numerically searchable property data)

Other Sources: EINECS\*\*

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

CM 1

CRN 32986-56-4 CMF C18 H37 N5 O9

Absolute stereochemistry.

CM 2

CRN 7664-93-9 CMF H2 O4 S

227 REFERENCES IN FILE CA (1907 TO DATE)
2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
228 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 144:376574

REFERENCE 2: 144:40605

REFERENCE 3: 144:31833

REFERENCE 4: 143:379779

REFERENCE 5: 143:205693

REFERENCE 6: 143:90148

REFERENCE 7: 142:469324

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8: 142:379031
REFERENCE
REFERENCE
             9:
                 142:348016
REFERENCE 10: 142:290622
L96 ANSWER 2 OF 4 REGISTRY COPYRIGHT 2006 ACS on STN
     32986-56-4 REGISTRY
RN
ED
     Entered STN: 16 Nov 1984
CN
     D-Streptamine, O-3-amino-3-deoxy-\alpha-D-glucopyranosyl-(1\rightarrow 6)-O-
     [2,6-diamino-2,3,6-trideoxy-\alpha-D-ribo-hexopyranosyl-(1\rightarrow4)]-2-
     deoxy- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Streptamine, 0-3-amino-3-deoxy-\alpha-D-glucopyranosyl-(1\rightarrow 4)-O-
     [2,6-diamino-2,3,6-trideoxy-\alpha-D-ribo-hexopyranosyl-(1\rightarrow6)]-2-
     deoxy-, D- (8CI)
OTHER NAMES:
     3'-Deoxykanamycin B
CN
CN
     Deoxykanamycin B
CN
     Nebramycin 6
CN
     Nebramycin factor 6
     Nebramycin VI
CN
     NF 6
CN
     NSC 180514
CN
     O-3-Amino-3-deoxy-\alpha-D-glucopyranosyl-(1\rightarrow 4)-O-[2,6-diamino-
CN
     2, 3, 6-trideoxy-\alpha-D-ribo-hexopyranosyl-(1\rightarrow 6)]-2-
     deoxystreptamine
CN
     Tobracin
     Tobradistin
CN
     Tobralex
CN
     Tobramaxin
CN
CN
     Tobramicin
CN
    Tobramycetin
CN
    Tobramycin
     Tobrex
CN
     STEREOSEARCH
FS
DR
     11098-01-4, 11111-45-8, 54330-95-9, 37321-13-4, 70322-33-7, 34337-51-4
MF
     C18 H37 N5 O9
CT
     COM
LC
     STN Files:
                   ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOSIS,
       BIOTECHNO, CA, CABA, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMLIST, CIN,
       CSCHEM, DDFU, DRUGU, EMBASE, HSDB*, IFICDB, IFIPAT, IFIUDB, IMSPATENTS,
       IPA, MEDLINE, MRCK*, NAPRALERT, PHAR, PIRA, PROMT, PROUSDDR, PS, RTECS*,
       SCISEARCH, SPECINFO, TOXCENTER, USAN, USPAT2, USPATFULL, VETU
          (*File contains numerically searchable property data)
                       EINECS**, WHO
          (**Enter CHEMLIST File for up-to-date regulatory information)
```

Absolute stereochemistry.

### \*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

4715 REFERENCES IN FILE CA (1907 TO DATE)
83 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
4725 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 144:456678 1: REFERENCE 2: 144:456193 REFERENCE 144:440079 3: REFERENCE 4: 144:425139 REFERENCE 144:419732 5: REFERENCE 144:408000 6: REFERENCE 7: 144:398353 REFERENCE 8: 144:397973 REFERENCE 9: 144:385541 REFERENCE 10: 144:382029

NEFERENCE 10: 144:302029

L96 ANSWER 3 OF 4 REGISTRY COPYRIGHT 2006 ACS on STN RN 75-65-0 REGISTRY

RN 75-65-0 REGISIRI

ED Entered STN: 16 Nov 1984

CN 2-Propanol, 2-methyl- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN tert-Butyl alcohol (8CI)

OTHER NAMES:

CN 1,1-Dimethylethanol CN 2-Methyl-2-propanol

CN t-Butanol

CN t-Butanol

CN tert-Butanol

CN Trimethylcarbinol

CN Trimethylmethanol

FS 3D CONCORD

MF C4 H10 O

CI COM

LC STN Files: AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN\*, BIOSIS, BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMINFORMRX, CHEMLIST,

CHEMSAFE, CIN, CSCHEM, CSNB, DDFU, DETHERM\*, DRUGU, EMBASE, ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT, ENCOMPPAT2, GMELIN\*, HSDB\*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, MSDS-OHS, PIRA, PROMT, PS, RTECS\*, SPECINFO, TOXCENTER, TULSA, ULIDAT, USPAT2, USPATFULL, VTB (\*File contains numerically searchable property data)

Other Sources: DSL\*\*, EINECS\*\*, TSCA\*\*

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

#### \*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

17366 REFERENCES IN FILE CA (1907 TO DATE)
310 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
17416 REFERENCES IN FILE CAPLUS (1907 TO DATE)
5 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 144:458328
REFERENCE 2: 144:457575
REFERENCE 3: 144:456604

REFERENCE 4: 144:456580

REFERENCE 5: 144:456420

REFERENCE 6: 144:456358

REFERENCE 7: 144:454943

REFERENCE 8: 144:450990

REFERENCE 9: 144:450736

REFERENCE 10: 144:450696

L96 ANSWER 4 OF 4 REGISTRY COPYRIGHT 2006 ACS on STN

RN **64-17-5** REGISTRY

ED Entered STN: 16 Nov 1984

CN Ethanol (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Ethyl alcohol (6CI, 7CI, 8CI)

OTHER NAMES:

CN 100C.NPA

CN AHD 2000

CN Alcare Hand Degermer

CN Alcohol

CN Alcohol anhydrous

CN Algrain

CN Anhydrol

```
Anhydrol PM 4085
CN
     CDA 19
CN
     CDA 19-200
CN
CN
     Desinfektol EL
     Duplicating Fluid 100C.NPA
CN
     Esumiru WK 88
CN
CN
     Ethicap
CN
     Ethyl hydrate
CN
     Ethyl hydroxide
CN
     Hinetoless
CN
     IMS 99
CN
     Infinity Pure
CN
     Jaysol
     Jaysol S
CN
CN
     Lux
CN
     Methylcarbinol
CN
     Molasses alcohol
     NSC 85228
CN
CN
     Potato alcohol
CN
     SDA 3A
     SDA 40-2
CN
     Sekundasprit
CN
     Sterillium Rub
CN
CN
     SY Fresh M
CN
    Synasol
CN
     Tecsol
CN
     Tecsol C
FS
     3D CONCORD
DR
     8000-16-6, 8024-45-1, 121182-78-3
ΜF
     C2 H6 O
CI
     COM
LC
     STN Files:
                  ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*, BIOSIS,
       BIOTECHNO, CA, CABA, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS,
       CHEMINFORMRX, CHEMLIST, CHEMSAFE, CIN, CSCHEM, CSNB, DDFU, DETHERM*,
       DRUGU, EMBASE, ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT, ENCOMPPAT2, GMELIN*,
       HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NAPRALERT,
       PIRA, PROMT, PS, RTECS*, SPECINFO, TOXCENTER, TULSA, ULIDAT, USAN,
       USPAT2, USPATFULL, VETU, VTB
         (*File contains numerically searchable property data)
                      DSL**, EINECS**, TSCA**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
{\rm H_3C-CH_2-OH}
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
          191910 REFERENCES IN FILE CA (1907 TO DATE)
            1635 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
          192841 REFERENCES IN FILE CAPLUS (1907 TO DATE)
              11 REFERENCES IN FILE CAOLD (PRIOR TO 1967)
REFERENCE
            1: 144:460046
REFERENCE
            2: 144:460014
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REFERENCE

3: 144:459999

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4: 144:459998
REFERENCE
REFERENCE
                144:459975
            5:
                144:459972
REFERENCE
            6:
               144:459913
REFERENCE
            7:
REFERENCE
            8:
                144:459705
REFERENCE
            9:
                144:459469
REFERENCE 10: 144:459213
=> d his
     (FILE 'HOME' ENTERED AT 10:35:25 ON 06 JUN 2006)
                SET COST OFF
     FILE 'REGISTRY' ENTERED AT 10:35:33 ON 06 JUN 2006
                E TOBRAMYCIN/CN
              1 S E3
L1
L2
             24 S C18H37N5O9/MF
L3
             24 S L2 AND OC5/ES AND 46.150.1/RID AND 3/NR
L4
              3 S L3 AND 2 3 6 TRIDEOXY AND 2 DEOXY AND 3 AMINO 3 DEOXY AND STR
L5
              2 S L4 NOT LABELED
              2 S L1, L5
L6
L7
             65 S 66007-88-3/CRN OR 32986-56-4/CRN
L8
             35 S L7 AND (PMS OR MXS OR IDS)/CI
1.9
             30 S L7 NOT L8
L10
             18 S L9 AND COMPD
T.11
             12 S L9 NOT L10
L12
             14 S L6, L11
L13
             10 S (METHANOL OR ETHANOL OR 1-PROPANOL OR 2-PROPANOL OR PROPANOL
     FILE 'HCAPLUS' ENTERED AT 10:40:36 ON 06 JUN 2006
L14
           4917 S L12
           4793 S TOBRAMYCIN?
L15
L16
             58 S TOBRAMICIN?
L17
           5447 S L14-L16
             55 S L17 AND L13
L18
L19
              4 S L12(L) PREP+NT/RL AND L18
L20
              2 S L19 NOT (137:30253 OR 102:119707)/DN
                E KWOK/AU
                E KWOK K/AU
L21
             16 S E3, E10, E11
L22
              5 S E30, E31
                E YANG/AU
L23
              4 S E3
                E YANG K/AU
T.24
            251 S E3
L25
             23 S E23
                E YANG KANG/AU
L26
            132 S E3, E8
                E ABRAXIS/PA,CS
                E AM PHARM/PA,CS
                E AM PHAR/PA, CS
                E A PHAR/PA,CS
                E AME PHAR/PA, CS
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E AMER PHAR/PA,CS
                 E AMERIC PHAR/PA,CS
                 E AMERICA PHAR/PA,CS
                 E AMERICAN PHAR/PA,CS
L27
               4 S E19-E22
L28
               0 S L17 AND L21-L27
L29
             113 S L17 (L) PREP+NT/RL
L30
             390 S L17 (L) PROC+NT/RL
L31
               9 S L29, L30 AND L18
L32
               5 S L31 NOT L19
L33
               1 S L32 AND 137:358002/DN
               3 S L20, L33
L34
     FILE 'REGISTRY' ENTERED AT 10:49:01 ON 06 JUN 2006
L35
               1 S 75-65-0
     FILE 'HCAPLUS' ENTERED AT 10:49:10 ON 06 JUN 2006
L36
               2 S L35 AND L17
L37
               3 S L34 AND L14-L34,L36
L38
           2193 S L17 AND (P AERUGINOSA? OR PROTEUS? OR P MIRABILIS OR M MORGAN
                 E STAPHYLOCCUS/CT
L39
               7 S E4+OLD, NT
L40
          39556 S E14+OLD, NT
          27095 S E21+OLD, NT
L41
L42
              36 S E23+OLD, NT OR E24+OLD, NT OR E25+OLD, NT
                 E PROVIDENCIA/CT
            938 S E3+OLD, NT
L43
L44
               2 S E15, E16
                 E CITROBACTER/CT
L45
           2742 S E3+OLD, NT
                 E KLEBSIELLA/CT
          10671 S E3+OLD, NT
L46
                 E ENTERBACTER/CT
                 E ENTEROBACTER/CT
L47
           6147 S E3+OLD, NT
                 E SERRATIA/CT
           6134 S E3+OLD, NT
T.48
                 E "E COLI"/CT
                 E ESCHERICHIA COLI/CT
         151013 S E3+OLD, NT
L49
                 E PROTEUS/CT
           6724 S E3+OLD, NT OR E5+OLD, NT
L50
L51
               2 S E7
                 E PROTEUS MIRABILIS/CT
L52
           2663 S E3+OLDNT
                 E MORGANELLA MORGANII/CT
L53
            905 S E3+OLD, NT
L54
            395 S E1+OLD, NT
L55
            437 S RETTGERI/CW
                 E PROVIDENCIA RETTGERI/CT
            426 S E3+OLD, NT
L56
                E PSEUDOMONAS RETTGERI/CT
L57
               1 S E3
L58
          22472 S AERUGINOSA/CW
                 E PSEUDOMONAS AERUGINOSA/CT
L59
          21201 S E3+OLD, NT
L60
          16976 S VULGARIS/CW
                 E PROTEUS VULGARIS/CT
           2281 S E3+OLD, NT
L61
L62
           2016 S L17 AND L39-L61
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L63
           2439 S L38, L62
                E SEPTICEMIA/CT
                E E3+ALL
           7766 S E4+OLD, NT
L64
L65
          28660 S E4/BI OR E6/BI OR E7/BI OR E9/BI OR E10/BI OR E11/BI OR E12/B
                E UNIARY TRACT INFECTION/CT
                E URINARY TRACT INFECTION/CT
                E E3+ALL
L66
            737 S E2
                E RESPIRATORY INFECTION/CT
                E E4+ALL
           1557 S E2
L67
                E SKIN INFECTION/CT
                E E3+ALL
           1298 S E2,E3
L68
                E SOFT TISSUE INFECTION/CT
                E E2+ALL
L69
            168 S E2 (L) INFECT? OR INFECTION?/CT (L) TISSUE(L)SOFT
                E BURN/CT
                E E3+ALL
L70
           7966 S E3+OLD, NT
                E PERITONITIS/CT
L71
            101 S E3+OLD, NT
                E E3+ALL
            743 S E2,E3
L72
                E CENTRAL NERVOUS SYSTEM INFECTION/CT
                E E3+ALL
            336 S E2, E3
L73
            211 S L17 AND L64-L73
L74
L75
            129 S L63 AND L74
L76
           2521 S L63, L74, L75
           2223 S L76 AND L14
L77
L78
              1 S L77 AND LYOPHIL?
L79
              2 S L76 AND LYOPHIL? .
L80
              5 S L78, L79, L34
L81
             31 S L76 AND LIQUID
L82
             15 S L76 AND AQUEOUS
             10 S L76 AND (FREEZ? OR FROZ? OR VACUUM?)
L83
L84
              5 S L81, L82 AND L83
                SEL DN 1 3 5
L85
              2 S L84 NOT E1-E3
L86
              6 S L80, L85
L87
              6 S L86 AND L14-L34,L36-L86
                E FREEZE DRYING/CT
L88
           6932 S E3+OLD, NT
                E SOLVENT/CT
          56978 S E64+OLD, NT
L89
L90
             11 S L17 AND L88
L91
              4 S L90 AND L89
L92
              3 S L90 AND L18
L93
              5 S L91, L92
L94
              4 S L93 NOT 136:391077/DN
L95
              9 S L87, L94 AND L14-L34, L36-L94
                SEL HIT RN
     FILE 'REGISTRY' ENTERED AT 11:11:02 ON 06 JUN 2006
L96
              4 S E1-E4
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FILE 'REGISTRY' ENTERED AT 11:11:23 ON 06 JUN 2006

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FILE COVERS 1907 - 6 Jun 2006 VOL 144 ISS 24 FILE LAST UPDATED: 5 Jun 2006 (20060605/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

#### => d 195 all hitstr tot

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L95 ANSWER 1 OF 9 HCAPLUS COPYRIGHT 2006 ACS on STN
```

AN 2005:369255 HCAPLUS

DN 142:397782

ED Entered STN: 29 Apr 2005

- TI Aqueous aerosol preparation as inhalants for drugs with unpleasant sensory characteristics containing nonionic surfactants and phospholipids
- IN Jauernig, Juergen; Lintz, Frank-Christophe; Keller, Manfred
- PA Pari GmbH, Germany
- SO PCT Int. Appl., 43 pp. CODEN: PIXXD2

DT Patent

LA German

IC ICM A61K0009-00

CC 63-6 (Pharmaceuticals)

FAN.CNT 2

r Alv. (	PATENT NO.						)	DATE			APPLICATION NO.						DATE			
PI		2005			A2 A3		20050428		WO 2004-EP11571						20041014					
		₩:	AE, CN, GE, LK,	AG, CO, GH, LR,	AL, CR, GM, LS,	AM, CU, HR, LT,	AT, CZ, HU, LU,	AU, DE, ID, LV,	AZ, DK, IL, MA,	DM, IN, MD,	DZ, IS, MG,	EC, JP, MK,	EE, KE, MN,	EG, KG, MW,	ES, KP, MX,	FI, KR, MZ,	GB, KZ, NA,	GD, LC, NI,		
		RW:	TJ, BW, AZ, EE, SI,	TM, GH, BY, ES, SK,	TN, GM, KG, FI, TR,	TR, KE, KZ, FR,	TT, LS, MD, GB,	PL, TZ, MW, RU, GR, CF,	UA, MZ, TJ, HU,	UG, NA, TM, IE,	US, SD, AT, IT,	UZ, SL, BE, LU,	VC, SZ, BG, MC,	VN, TZ, CH, NL,	YU, UG, CY, PL,	ZA, ZM, CZ, PT,	ZM, ZW, DE, RO,	ZW AM, DK, SE,		
PRAI	US	SN, TD, TG E 10347994 S 2005244339 E 2003-10347994				A1 A1 A		2005 2005 2003	1103		DE 2003-10347994 US 2005-106999						20031015 20050414			

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WO 2004-EP11571
                               20041014
                         Α2
CLASS
 PATENT NO.
                CLASS PATENT FAMILY CLASSIFICATION CODES
 _____
                ____
                       _________
 WO 2005037246
                ICM
                       A61K0009-00
                IPCI
                       A61K0009-00 [ICM,7]; A61K0009-107 [ICS,7]; A61K0009-00
                       [ICS,7]; A61K0009-12 [ICS,7]
                IPCR
                       A61K0009-107 [I,A]; A61K0009-107 [I,C*]; A61K0009-127
                       [N,A]; A61K0009-127 [N,C*]; A61K0009-19 [N,A];
                       A61K0009-19 [N,C*]
                ECLA
                       A61K009/00M20B; A61K009/107D
 DE 10347994
                IPCI
                       A61K0047-18 [ICM,7]; A61K0047-16 [ICM,7,C*];
                       A61P0043-00 [ICS,7]
                       A61K0009-107 [I,A]; A61K0009-107 [I,C*]; A61K0009-127
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                       [N,A]; A61K0009-127 [N,C^*]; A61K0009-19 [N,A];
                       A61K0009-19 [N,C*]
                ECLA
                       A61K009/00M20B
 US 2005244339
                       A61L0009-04 [ICM,7]
                IPCI
                IPCR
                       A61K0009-107 [I,A]; A61K0009-107 [I,C*]; A61K0009-127
                       [N,A]; A61K0009-127 [N,C*]; A61K0009-19 [N,A];
                       A61K0009-19 [N,C*]
                NCL
                       424/045.000
                ECLA
                       A61K009/00M20B; A61K009/107D
AB
    Disclosed are sterile aqueous prepns. that are to be inhaled as an aerosol and
    contain an active substance, a nonionic surfactant, and a phospholipid.
    Said prepns. are suitable for administering poorly soluble active substances
    by way of inhalation in the form of colloidal solns. and can also be used
    for administering bad-tasting active substances that irritate the mucus
    and cause cough or bronchoconstrictions. The inventive prepns. can be
    nebulized by means of conventional devices and are preferably used in
    pediatrics. Thus a 1000 mL aqueous formulation contained (g): Budesonide 0.2;
    Tyloxapol 10.0; DMPC 5.0; sodium chloride 8.4; citric acid/sodium acetate
    to pH 4.4.
ST
    inhalant budesonide aq aerosol phospholipid nonionic surfactant Tyloxapol
TT
    Drug delivery systems
        (aerosols; aqueous aerosol preparation as inhalants for drugs with
unpleasant
        sensory characteristics containing nonionic surfactants and phospholipids)
    Tumor necrosis factors
TT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (antibodies to; aqueous aerosol preparation as inhalants for drugs with
        unpleasant sensory characteristics containing nonionic surfactants and
       phospholipids)
IT
    Anti-infective agents
    Atomizing (spraying)
    Cholinergic antagonists
    Cough
    Cytotoxic agents
       Freeze drying
    Human
    Immunomodulators
    Mammalia
    Osmolality
    Packaging materials
    Particle size
    Sols
    Solubility
    Sonication
    Sterilization and Disinfection
    Surface tension
```

Viscosity pH

(aqueous aerosol preparation as inhalants for drugs with unpleasant sensory characteristics containing nonionic surfactants and phospholipids)

IT Corticosteroids, biological studies

Lecithins

Phospholipids, biological studies

Polyoxyalkylenes, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(aqueous aerosol preparation as inhalants for drugs with unpleasant sensory characteristics containing nonionic surfactants and phospholipids)

IT Bronchi

 $\hbox{(bronchoconstriction; aqueous aerosol preparation as inhalants for drugs} \\$  with

unpleasant sensory characteristics containing nonionic surfactants and phospholipids)

IT Development, mammalian postnatal

(child; aqueous aerosol preparation as inhalants for drugs with unpleasant sensory characteristics containing nonionic surfactants and phospholipids)

IT Solvents

TΤ

IT

IT

(cosolvents; aqueous aerosol preparation as inhalants for drugs with unpleasant

sensory characteristics containing nonionic surfactants and phospholipids)
IT Development, mammalian postnatal

(infant; aqueous aerosol preparation as inhalants for drugs with unpleasant sensory characteristics containing nonionic surfactants and phospholipids)
Drug delivery systems

(inhalants; aqueous aerosol preparation as inhalants for drugs with

sensory characteristics containing nonionic surfactants and phospholipids) IT Medical goods

(inhalers; aqueous aerosol preparation as inhalants for drugs with unpleasant

sensory characteristics containing nonionic surfactants and phospholipids) IT Anesthetics

(local; aqueous aerosol preparation as inhalants for drugs with unpleasant sensory characteristics containing nonionic surfactants and phospholipids) Surfactants

(nonionic; aqueous aerosol preparation as inhalants for drugs with unpleasant

sensory characteristics containing nonionic surfactants and phospholipids) IT Homogenization

(pressure; aqueous aerosol preparation as inhalants for drugs with unpleasant

sensory characteristics containing nonionic surfactants and phospholipids) IT Inflammation

Respiratory system, disease

(sinusitis; aqueous aerosol preparation as inhalants for drugs with unpleasant

sensory characteristics containing nonionic surfactants and phospholipids) IT Filtration

(sterile; aqueous aerosol preparation as inhalants for drugs with unpleasant sensory characteristics containing nonionic surfactants and phospholipids) Antibodies and Immunoglobulins

RL: BSU (Biological study, unclassified); BIOL (Biological study) (to  $TNF\alpha$ ; aqueous aerosol preparation as inhalants for drugs with unpleasant sensory characteristics containing nonionic surfactants and phospholipids)

IT Adrenoceptor antagonists

 $(\beta-;$  aqueous aerosol preparation as inhalants for drugs with unpleasant

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sensory characteristics containing nonionic surfactants and phospholipids)
     137-58-6, Lidocaine 32986-56-4, Tobramycin
TΤ
     51333-22-3, Budesonide
                              79217-60-0, Cyclosporin
     RL: PEP (Physical, engineering or chemical process); PYP
     (Physical process); THU (Therapeutic use); BIOL (Biological study);
     PROC (Process); USES (Uses)
        (aqueous aerosol preparation as inhalants for drugs with unpleasant sensory
        characteristics containing nonionic surfactants and phospholipids)
     56-81-5, Glycerol, biological studies
IT
                                            57-55-6, Propylene glycol,
     biological studies
                         58-95-7, Vitamin E acetate
                                                       59-05-2, Methotrexate
     70-18-8, Glutathione, biological studies
                                               114-07-8, Erythromycin
     139-33-3
                154-93-8, Carmustine
                                     302-79-4, Tretinoin
                                                            443-48-1,
     Metronidazol
                    446-86-6, Azathioprin
                                           616-91-1, Acetylcysteine
     1397-89-3, Amphotericin B
                                1400-61-9, Nystatin 2644-64-6, DPPC
     3056-17-5, Stavudine 3385-03-3, Flunisolide
                                                     4419-39-0, Beclomethasone
     4539-70-2, DSPC
                       6493-05-6
                                                        13010-47-4, Lomustine
                                   7681-93-8, Natamycin
                             18656-38-7, DMPC
     15663-27-1, Cisplatin
                                                23593-75-1, Clotrimazole
     25301-02-4, Tyloxapol
                             25322-68-3, Polyethylene glycol
                                                              30516-87-1,
                  33069-62-4, Taxol
     Zidovudine
                                      33419-42-0, Etoposide
                                                              34391-04-3.
     Levalbuterol
                    59277-89-3, Aciclovir
                                            60205-81-4, Ipratropium
     60628-96-8, Bifonazole
                              65277-42-1, Ketoconazole
                                                         69655-05-6, Didanosine
     73573-87-2, Formoterol
                              81103-11-9, Clarithromycin
                                                           83905-01-5,
     Azithromycin
                    84625-61-6, Itraconazole
                                               85721-33-1, Ciprofloxacin
     86386-73-4, Fluconazole 89365-50-4, Salmeterol
                                                        90566-53-3, Fluticasone
     99571-64-9, Oxitropium 104227-87-4, Famciclovir
                                                         104987-11-3,
     Tacrolimus
                  105102-22-5, Mometasone 124832-26-4, Valaciclovir
     126544-47-6, Ciclesonide
                               127779-20-8, Saguinavir
                                                         134678-17-4,
                 151096-09-2, Moxifloxacin 155213-67-5, Ritonavir
     Lamivudine
     186691-13-4, Tiotropium
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (aqueous aerosol preparation as inhalants for drugs with unpleasant sensory
        characteristics containing nonionic surfactants and phospholipids)
     9015-82-1
IT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (inhibitors; aqueous aerosol preparation as inhalants for drugs with
unpleasant
        sensory characteristics containing nonionic surfactants and phospholipids)
     32986-56-4, Tobramycin
     RL: PEP (Physical, engineering or chemical process); PYP
     (Physical process); THU (Therapeutic use); BIOL (Biological study);
     PROC (Process); USES (Uses)
        (aqueous aerosol preparation as inhalants for drugs with unpleasant sensory
        characteristics containing nonionic surfactants and phospholipids)
RN
     32986-56-4 HCAPLUS
CN
     D-Streptamine, O-3-amino-3-deoxy-\alpha-D-glucopyranosyl-(1\rightarrow6)-O-
     [2,6-diamino-2,3,6-trideoxy-\alpha-D-ribo-hexopyranosyl-(1\rightarrow4)]-2-
     deoxy- (9CI) (CA INDEX NAME)
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L95
     ANSWER 2 OF 9 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     2003:576810 HCAPLUS
DN
     139:306572
ED
     Entered STN: 29 Jul 2003
     Improvement on crystallization process of tobramycin
ΤI
ΑU
     Wang, Hong; Yu, Shujuan; Gao, Dawei
CS
     Department of Biological Engineering, South China University of
     Technology, Canton, 510640, Peop. Rep. China
SO
     Zhongguo Yiyao Gongye Zazhi (2003), 34(1), 13-14
     CODEN: ZYGZEA; ISSN: 1001-8255
PB
     Zhongguo Yiyao Gongye Zazhi Bianjibu
DΤ
     Journal
LA
     Chinese
CC
     16-1 (Fermentation and Bioindustrial Chemistry)
AB
     The crystallization process of tobramycin in the Streptomyces
     tenebrarius fermentation broth was improved to increase the crystallization
rate as well
     as the titer by adjusting several factors such as seeding, volume of
     ethanol, stirring rate, and washing solvent.
ST
     tobramycin crystn improvement
IT
     Fermentation
        (broth; improvement on crystallization process of tobramycin in
        fermentation broth)
IT
    Agitation (mechanical)
    Crystallization
    Streptomyces tenebrarius
        (improvement on crystallization process of tobramycin in fermentation
        broth)
IT
     64-17-5, Ethanol, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (improvement on crystallization process of tobramycin in fermentation
        broth)
     32986-56-4P, Tobramycin
IΤ
     RL: PUR (Purification or recovery); PREP (Preparation)
        (improvement on crystallization process of tobramycin in fermentation
        broth)
ΙT
     64-17-5, Ethanol, biological studies
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (improvement on crystallization process of tobramycin in fermentation
        broth)
RN
     64-17-5 HCAPLUS
CN
    Ethanol (9CI) (CA INDEX NAME)
```

H3C-CH2-OH

# IT 32986-56-4P, Tobramycin

RL: PUR (Purification or recovery); PREP (Preparation) (improvement on crystallization process of tobramycin in fermentation broth)

RN 32986-56-4 HCAPLUS

CN D-Streptamine, O-3-amino-3-deoxy- $\alpha$ -D-glucopyranosyl- $(1\rightarrow 6)$ -O-[2,6-diamino-2,3,6-trideoxy- $\alpha$ -D-ribo-hexopyranosyl- $(1\rightarrow 4)$ ]-2-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L95 ANSWER 3 OF 9 HCAPLUS COPYRIGHT 2006 ACS on STN

AN 2002:754995 HCAPLUS

DN 137:268473

ED Entered STN: 04 Oct 2002

TI Porous drug matrices and methods of manufacture thereof

IN Straub, Julie; Altreuter, David; Bernstein, Howard; Chickering, Donald E.; Khattak, Sarwat; Randall, Greg

PA Acusphere Inc., USA

SO U.S. Pat. Appl. Publ., 20 pp., Cont.-in-part of U.S. 6,395,300. CODEN: USXXCO

DT Patent

LA English

IC ICM A61K0009-14 ICS A61K0009-50

INCL 424499000

CC 63-6 (Pharmaceuticals)

FAN CNT 2

FAN.	JNT	2																	
	PAT	CENT	NO.			KIND		DATE			APPLICATION NO.						DATE		
							-												
ΡI	US	2002	1420	50		A1		20021003			US	2002-	-5392	20020122					
	US	6395	300			В1		20020528			US	1999-	-4334	19991104					
	EΡ	1642	572			A1		2006	20060405			2005-	-2719	20000525					
		R:	AT,	BE,	CH,	DE,	DK	, ES,	FR,	GB,	GF	R, IT,	LI,	LU,	NL,	SE,	MC,	PT,	
			ΙE,	FI,	CY														
	US	6645528				В1		2003	20031111			2000-	-6944	20001023					
	US	6932	В1		2005	0823		US	2000-	-7060	45		20	0001	103				
	za	A 2001010347						2003	0730		ZA	2001-	-1034	7		20	0011	218	
	US	S 2005048116						20050303			US 2004-924642					20040824			
	US	JS 2005058710						2005	0317		US	2004-	-9288	86		20	0040	B27	
PRAI	US	1999	-136	323P		P		1999	0527										
	US	1999	-158	659P		P		1999	1008										

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US 1999-433486
                          A2
                                 19991104
                         Р
     US 2000-186310P
                                 20000302
     EP 2000-939365
                          Α3
                                 20000525
     US 2002-53929
                          А3
                                 20020122
CLASS
 PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
                 ____
 US 2002142050
                 ICM
                        A61K0009-14
                 ICS
                        A61K0009-50
                 INCL
                        424499000
                 IPCI
                        A61K0009-14 [ICM, 7]; A61K0009-50 [ICS, 7]
                 IPCR
                        A61K0009-16 [I,A]; A61K0009-16 [I,C*]
                 NCL
                        424/499.000
                        A61K009/16P4; A61K009/16P2
                 ECLA
 US 6395300
                        A61K0009-14 [ICM,7]; A61K0047-02 [ICS,7]; B29B0009-00
                 IPCI
                        [ICS, 7]
                 IPCR
                        A61K0009-16 [I,A]; A61K0009-16 [I,C*]
                 NCL
                        424/489.000; 264/005.000; 977/906.000
                 ECLA
                        A61K009/16P4; A61K009/16P2
 EP 1642572
                        A61K0009-16 [I,A]
                 IPCI
 US 6645528
                 IPCI
                        A61K0009-14 [ICM, 7]
                 IPCR
                        A61K0009-16 [I,A]; A61K0009-16 [I,C*]
                 NCL
                        424/489.000; 514/951.000; 977/923.000
                 ECLA
                        A61K009/16H2; A61K009/16H6B; A61K009/16H4B;
                        A61K009/16P4; A61K009/16P2
 US 6932983
                 IPCI
                        A61K0009-16 [ICM, 7]
                 IPCR
                        A61K0009-16 [I,A]; A61K0009-16 [I,C*]
                 NCL
                        424/489.000; 424/400.000
                 ECLA
                        A61K009/16H2; A61K009/16H4B; A61K009/16H6B;
                        A61K009/16P2; A61K009/16P4
 ZA 2001010347
                 IPCI
                        A61K [ICM, 7]
 US 2005048116
                 IPCI
                        A61K0009-26 [ICM, 7]; A61K0009-14 [ICS, 7]
                 IPCR
                        A61K0009-16 [I,A]; A61K0009-16 [I,C*]
                 NCL
                        424/469.000
                 ECLA
                        A61K009/16H2; A61K009/16H4B; A61K009/16H6B;
                        A61K009/16P2; A61K009/16P4
 US 2005058710
                 IPCI
                        A61K0009-26 [ICM, 7]; A61K0009-14 [ICS, 7]
                 IPCR
                        A61K0009-16 [I,A]; A61K0009-16 [I,C*]
                 NCL
                        424/469.000
                 ECLA
                        A61K009/16H2; A61K009/16H4B; A61K009/16H6B;
                        A61K009/16P2; A61K009/16P4
AB
     Drugs, especially low aqueous solubility drugs, are provided in a porous
matrix form,
     preferably microparticles, which enhances dissoln. of the drug in agueous
     media. The drug matrixes preferably are made using a process that
     includes (i) dissolving a drug, preferably a drug having low aqueous
solubility, in
     a volatile solvent to form a drug solution, (ii) combining at least one pore
     forming agent with the drug solution to form an emulsion, suspension, or
     second solution and hydrophilic or hydrophobic excipients that stabilize the
     drug and inhibit crystallization, and (iii) removing the volatile solvent and
pore
     forming agent from the emulsion, suspension, or second solution to yield the
     porous matrix of drug. Hydrophobic or hydrophilic excipients may be
     selected to stabilize the drug in crystalline form by inhibiting crystal growth
     or to stabilize the drug in amorphous form by preventing crystallization The
pore
     forming agent can be either a volatile liquid that is immiscible with the
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drug solvent or a volatile solid compound, preferably a volatile salt. In a preferred embodiment, spray drying is used to remove the solvents and the

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pore forming agent. The resulting porous matrix has a faster rate of
dissoln. following administration to a patient, as compared to non-porous
matrix forms of the drug. In a preferred embodiment, microparticles of
the porous drug matrix are reconstituted with an aqueous medium and
administered parenterally, or processed using standard techniques into tablets
or capsules for oral administration. Thus, 5.46 g of PEG 8000, 0.545 g of
prednisone, and 0.055 g of Span 40 were dissolved in 182 mL of methylene
chloride. A solution of 3.27 g of ammonium bicarbonate in 18.2 mL of water
was added to the organic solution (phase ratio 1:10) and homogenized for 5 min
at 16,000 RPM. The resulting emulsion was spray dried on a benchtop spray
dryer using an air-atomizing nozzle and nitrogen as the drying gas.
porous drug matrix microparticle prednisone bicarbonate
Drug delivery systems
   (buccal; porous drug matrixes and methods of manufacture thereof)
Estrogens
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
   (conjugated; porous drug matrixes and methods of manufacture thereof)
Drying
   (fluid bed; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (inhalants; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (injections, i.m.; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (injections, i.v.; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (injections, s.c.; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (microparticles; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (nasal; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (ophthalmic; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (oral; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (parenterals; porous drug matrixes and methods of manufacture thereof)
Dissolution
  Freeze drying
Preservatives
  Solvents
   (porous drug matrixes and methods of manufacture thereof)
Amino acids, biological studies
Carbohydrates, biological studies
Granulocyte colony-stimulating factor receptors
Interferons
Interleukins
Lecithins
Polymers, biological studies
Polyoxyalkylenes, biological studies
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
   (porous drug matrixes and methods of manufacture thereof)
Crystallization
   (prevention of; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (rectal; porous drug matrixes and methods of manufacture thereof)
Drug delivery systems
   (sublingual; porous drug matrixes and methods of manufacture thereof)
Drying
   (vacuum; porous drug matrixes and methods of manufacture thereof)
```

IT Drug delivery systems

TT

(vaginal; porous drug matrixes and methods of manufacture thereof) 50-28-2, Estradiol, biological studies 50-35-1, Thalidomide 53-03-2, Prednisone 55-98-1, Busulfan Verapamil 57-63-6, Ethinyl 58-61-7, Adenosine, biological studies 59-92-7, Levodopa, estradiol biological studies 67-78-7 67-97-0, Vitamin D3 71-58-9, Medroxyprogesterone acetate 75-64-9, Erbumine, biological studies 77-36-1, Chlorthalidone 89-57-6, Mesalamine 126-07-8, Griseofulvin 128-13-2, Ursodiol 298-46-4, Carbamazepine 302-79-4, Tretinoin 321-64-2, Tacrine 363-24-6, Dinoprostone 437-38-7, Fentanyl 439-14-5, Diazepam 443-48-1, Metronidazole 518-28-5, Podofilox 631-61-8, Ammonium acetate 657-24-9, Metformin 745-65-3, Alprostadil 846-49-1, Lorazepam 1066-33-7, Ammonium bicarbonate 1863-63-4, Ammonium benzoate 1951-25-3, Amiodarone 3239-44-9, Dexfenfluramine 4759-48-2, Isotretinoin 5534-09-8, Beclomethasone dipropionate 5593-20-4, Betamethasone dipropionate 9002-68-0, Follitropin 9002-72-6, Growth hormone 9005-65-6, Tween 80 9007-12-9, Calcitonin 9041-93-4, Bleomycin sulfate 10238-21-8, Glyburide 11096-26-7, 12125-02-9, Ammonium chloride, biological studies Erythropoietin 12629-01-5, Somatropin 12633-72-6, Amphotericin 13311-84-7, Flutamide 15307-79-6, Diclofenac sodium 15307-86-5, Diclofenac 15687-27-1, Ibuprofen 18559-94-9, Albuterol 20830-75-5, Digoxin 21256-18-8, Oxaprozin 21829-25-4, Nifedipine 22204-53-1, Naproxen 25322-68-3, Polyethylene glycol 26266-57-9, Span 40 27203-92-5, Tramadol 28860-95-9, Carbidopa 28981-97-7, Alprazolam. 29094-61-9, Glipizide 30516-87-1, Zidovudine **32986-56-4**, Tobramycin 33069-62-4, Paclitaxel 34911-55-2, Bupropion 36505-84-7, Buspirone 40391-99-9 41340-25-4, Etodolac 41575-94-4, Carboplatin 42399-41-7, Diltiazem 42924-53-8, Nabumetone 51333-22-3, Budesonide 51773-92-3, Mefloquine hydrochloride 54143-55-4, Flecainide 54527-84-3, Nicardipine hydrochloride 54910-89-3, Fluoxetine 54965-21-8, 55268-75-2, Cefuroxime Albendazole 54965-24-1, Tamoxifen citrate 56124-62-0, Valrubicin 56180-94-0, Acarbose 60142-96-3, Gabapentin 60205-81-4, Ipratropium. 63659-18-7, Betaxolol 65277-42-1, Ketoconazole 66085-59-4, Nimodipine 66376-36-1, Alendronate 66852-54-8, Halobetasol propionate 68693-11-8, Modafinil 69655-05-6, Didanosine 70476-82-3, Mitoxantrone hydrochloride 72432-03-2, Miglitol 72509-76-3, Felodipine 72558-82-8, Ceftazidime 72956-09-3, Carvedilol 73384-59-5, Ceftriaxone 73590-58-6, Omeprazole 75330-75-5, Lovastatin 75695-93-1, Isradipine 76095-16-4, Enalapril 75847-73-3, Enalapril 76547-98-3, Lisinopril maleate 76824-35-6, Famotidine Nizatidine 77883-43-3, Doxazosin mesylate 78246-49-8, Paroxetine hydrochloride 78628-80-5, Terbinafine hydrochloride 78755-81-4, Flumazenil 79517-01-4, Octreotide acetate 79559-97-0, Sertraline hydrochloride 79794-75-5, Loratadine 79902-63-9, Simvastatin 80274-67-5, Metoprolol fumarate 81098-60-4, Cisapride 81103-11-9, Clarithromycin 82410-32-0, Ganciclovir 82752-99-6, Nefazodone hydrochloride 82834-16-0, Perindopril 83799-24-0, Fexofenadine 83905-01-5, Azithromycin 83919-23-7, Mometasone furoate 84625-61-6, Itraconazole 86386-73-4, Fluconazole 86541-74-4, Benazepril hydrochloride 86541-75-5, Benazepril 87679-37-6, Trandolapril 89778-27-8, Toremifene citrate 90566-53-3, Fluticasone 91161-71-6, Terbinafine 91421-42-0, Rubitecan 93413-69-5, Venlafaxine 93957-54-1, Fluvastatin 95058-81-4, Gemcitabine 95233-18-4, Atovaquone 97048-13-0, Urofollitropin 97322-87-7, Troglitazone 98048-97-6, Fosinopril 98079-52-8, Lomefloxacin hydrochloride 98319-26-7, 99011-02-6, Imiquimod Finasteride 99294-93-6, Zolpidem tartrate 100286-90-6, Irinotecan hydrochloride 100986-85-4, Levofloxacin 103577-45-3, Lansoprazole 103628-48-4, Sumatriptan succinate 103775-10-6, Moexipril 104227-87-4, Famciclovir 104632-25-9,

Pramipexole dihydrochloride 106266-06-2, Risperidone 106392-12-5. Pluronic f127 106463-17-6, Tamsulosin hydrochloride 106685-40-9, Adapalene 107753-78-6, Zafirlukast 109889-09-0, Granisetron 110871-86-8, Sparfloxacin 111470-99-6, Amlodipine besylate 111974-72-2, Quetiapine fumarate 112809-51-5, Letrozole Olopatadine 114798-26-4, Losartan 114977-28-5, Docetaxel 115956-12-2, Dolasetron 120014-06-4, Donepezil 124832-26-4, Valacyclovir 127779-20-8, Saguinavir 131918-61-1, Paricalcitol 132539-06-1, Olanzapine 134308-13-7, Tolcapone 134678-17-4, Lamivudine 137862-53-4, Valsartan 140678-14-4, Mangafodipir trisodium 142373-60-2, Tirofiban hydrochloride 144701-48-4, Telmisartan 145040-37-5, Candesartan cilexetil 147059-72-1, Trovafloxacin 147245-92-9, Glatiramer acetate 150378-17-9, Indinavir 154248-97-2, Imiglucerase 154598-52-4, Efavirenz 155141-29-0, Rosiglitazone maleate 155213-67-5, Ritonavir 158966-92-8, Montelukast 159989-65-8, Nelfinavir mesylate 162011-90-7, Rofecoxib 161814-49-9, Amprenavir 169590-42-5, Celecoxib 171599-83-0, Sildenafil citrate 260779-88-2, Cisapride monohydrate 679809-58-6, Enoxaparin sodium RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (porous drug matrixes and methods of manufacture thereof) ΙT 32986-56-4, Tobramycin RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (porous drug matrixes and methods of manufacture thereof) RN 32986-56-4 HCAPLUS CN D-Streptamine, O-3-amino-3-deoxy- $\alpha$ -D-glucopyranosyl- $(1\rightarrow 6)$ -O-[2,6-diamino-2,3,6-trideoxy- $\alpha$ -D-ribo-hexopyranosyl-(1 $\rightarrow$ 4)]-2deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L95

ANSWER 4 OF 9 HCAPLUS COPYRIGHT 2006 ACS on STN

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ΑN
     2002:588609 HCAPLUS
DN
     138:205270
     Entered STN: 08 Aug 2002
ED
     Research on influential factors and optimization in the process of
TI
     crystallization of Tobramycin
     Wang, Hong; Luo, Wenbo; Yu, Shujuan; Gao, Dawei
ΑU
CS
     Food and Biological Engineering College, SCUT, Canton, 510640, Peop. Rep.
     China
SO
     Zhongguo Kangshengsu Zazhi (2002), 27(4), 221-223
     CODEN: ZKZAEY; ISSN: 1001-8689
PB
     Zhongguo Kangshengsu Zazhishe
DT
     Journal
LA
     Chinese
CC
     33-7 (Carbohydrates)
AB
     Tobramycin was crystallized from its solution in ethanol (at ratio of
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1:10) at  $25^{\circ}$  for 10 h. The factors such as ethanol addition volume, temperature, time, and stirring rate in the process for crystallization of **Tobramycin** with the final yield, and titer indexes were analyzed by orthogonal method.

ST tobramycin crystn factor

IT Crystallization

(process for crystallization of Tobramycin)

IT 32986-56-4P, Tobramycin

RL: PNU (Preparation, unclassified); PUR (Purification or recovery); PREP (Preparation)

(process for crystallization of)

IT **64-17-5**, Ethanol, uses

RL: NUU (Other use, unclassified); USES (Uses) (process for crystallization of Tobramycin)

IT 32986-56-4P, Tobramycin

RL: PNU (Preparation, unclassified); PUR (Purification or recovery); PREP (Preparation)

(process for crystallization of)

RN 32986-56-4 HCAPLUS

CN D-Streptamine, O-3-amino-3-deoxy- $\alpha$ -D-glucopyranosyl- $(1\rightarrow 6)$ -O-[2,6-diamino-2,3,6-trideoxy- $\alpha$ -D-ribo-hexopyranosyl- $(1\rightarrow 4)$ ]-2-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

IT **64-17-5**, Ethanol, uses

RL: NUU (Other use, unclassified); USES (Uses) (process for crystallization of Tobramycin)

RN 64-17-5 HCAPLUS

CN Ethanol (9CI) (CA INDEX NAME)

H3C-CH2-OH

L95 ANSWER 5 OF 9 HCAPLUS COPYRIGHT 2006 ACS on STN

AN 2002:316298 HCAPLUS

DN 137:358002

ED Entered STN: 28 Apr 2002

TI Freeze-drying of tert-butanol/water cosolvent systems: a case report on formation of a friable freeze-dried powder of tobramycin sulfate

AU Wittaya-Areekul, Sakchai; Needham, Gregory F.; Milton, Nathaniel; Roy, Michael L.; Nail, Steven L.

CS Department of Industrial and Physical Pharmacy, School of Pharmacy, Purdue University, West Lafayette, IN, 47907, USA

SO Journal of Pharmaceutical Sciences (2002), 91(4), 1147-1155

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CODEN: JPMSAE; ISSN: 0022-3549
PB
    Wiley-Liss, Inc.
DT
     Journal
LA
     English
CC
     63-6 (Pharmaceuticals)
AΒ
    A case study is presented in which a tert-butanol (TBA)/water cosolvent
     system was found to be a useful means of producing freeze-dried
     tobramycin sulfate that readily forms a loose powder upon
     agitation in a specialized application in which a critical quality attribute
     is the ability to pour the sterile powder from the vial. Both formulation
     and processing variables are important in achieving acceptable phys.
     properties of the cake as well as minimizing residual TBA levels.
     Liquid/liquid phase separation was observed above critical concns. of both
drug and TBA,
     resulting in a two-layered lyophilized cake with unacceptable appearance,
    phys. properties, and residual TBA levels. However, the choice of
     tobramycin sulfate and TBA concns. in the single-phase region of
     the phase diagram resulted in a lyophilized solid that can readily be
    poured from vials. Crystallization of TBA before drying is critical to
achieving
    adequately low residual TBA levels, and this is reflected in the effect of
     thermal history of freezing on residual TBA levels, where rapid freezing
     results in incomplete crystallization of TBA and relatively high levels of
     residual solvent. Annealing at a temperature above T'g of the system after an
     initial freezing step significantly reduces the level of residual TBA.
     Secondary drying, even at increased temperature and for extended times, is not
    an effective method of reducing residual TBA levels.
ST
    freeze dried powder tobramycin butanol cosolvent
IT
    Crystallization
      Freeze drying
     Friability
     Phase separation
     Phase transition
        (butanol/water cosolvent system for producing freeze-dried
        tobramycin sulfate)
TΤ
    Solvents
        (cosolvents; butanol/water cosolvent system for producing freeze-dried
        tobramycin sulfate)
TT
    Drug delivery systems
        (powders; butanol/water cosolvent system for producing freeze-dried
        tobramycin sulfate)
IT
    75-65-0, tert-Butanol, uses
                                 7732-18-5, Water, uses
    RL: MOA (Modifier or additive use); USES (Uses)
        (butanol/water cosolvent system for producing freeze-dried
        tobramycin sulfate)
IT
    49842-07-1, Tobramycin sulfate
    RL: PEP (Physical, engineering or chemical process); PYP
     (Physical process); THU (Therapeutic use); BIOL (Biological study);
    PROC (Process); USES (Uses)
        (butanol/water cosolvent system for producing freeze-dried
        tobramycin sulfate)
RE.CNT
              THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD
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     75-65-0, tert-Butanol, uses
     RL: MOA (Modifier or additive use); USES (Uses)
        (butanol/water cosolvent system for producing freeze-dried
        tobramycin sulfate)
RN
     75-65-0 HCAPLUS
CN
     2-Propanol, 2-methyl- (9CI) (CA INDEX NAME)
    OH
H3C-C-CH3
    CH<sub>3</sub>
IT
     49842-07-1, Tobramycin sulfate
     RL: PEP (Physical, engineering or chemical process); PYP
     (Physical process); THU (Therapeutic use); BIOL (Biological study);
     PROC (Process); USES (Uses)
        (butanol/water cosolvent system for producing freeze-dried
        tobramycin sulfate)
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D-Streptamine, O-3-amino-3-deoxy- $\alpha$ -D-glucopyranosyl-(1 $\rightarrow$ 6)-O-

[2,6-diamino-2,3,6-trideoxy- $\alpha$ -D-ribo-hexopyranosyl-(1 $\rightarrow$ 4)]-2-

deoxy-, sulfate (2:5) (salt) (9CI) (CA INDEX NAME)

CM 1

RN

CN

CRN 32986-56-4 CMF C18 H37 N5 O9

49842-07-1 HCAPLUS

Absolute stereochemistry.

CM 2

CRN 7664-93-9 CMF H2 O4 S

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L95 ANSWER 6 OF 9 HCAPLUS COPYRIGHT 2006 ACS on STN
     2000:861473 HCAPLUS
DN
     134:32972
ED
     Entered STN: 08 Dec 2000
     Porous drug matrixes containing polymers and sugars and methods of their
     Straub, Julie; Bernstein, Howard; Chickering, Donald E., III; Khatak,
     Sarwat; Randall, Greg
PA
    Acusphere, Inc., USA
     PCT Int. Appl., 45 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
     ICM A61K0009-16
IC
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 1
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                        KIND
                                DATE
                                          APPLICATION NO.
                                                                  DATE
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         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
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PRAI US 1999-136323P P 19990527
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CLASS
                 CLASS PATENT FAMILY CLASSIFICATION CODES
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                         A61K0009-16
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                         A61K0009-16 [ICM,7]
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                 ECLA
                         A61K009/16H4B; A61K009/16H6B; A61K009/16P2;
                         A61K009/16P4
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                         A61K0009-14 [ICM,7]; A61K0047-02 [ICS,7]; B29B0009-00
                         [ICS, 7]
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                         A61K0009-16 [I,A]; A61K0009-16 [I,C*]
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                         424/489.000; 264/005.000; 977/906.000
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                 IPCR
                        A61K0009-16 [I,A]; A61K0009-16 [I,C*]
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                        A61K009/16P4
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                         [ICS,7]; A61K0009-10 [ICS,7]; A61K0009-20 [ICS,7];
                         A61K0009-48 [ICS,7]; A61K0047-02 [ICS,7]; A61K0047-12
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                        A61K0009-48 [ICM,7]; A61K0009-20 [ICS,7]; A61K0009-14
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                        A61K0009-16 [I,A]; A61K0009-16 [I,C*]
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                        424/452.000
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                        A61K009/16H4B; A61K009/16H6B; A61K009/16H2;
                        A61K009/16P4
 NO 2001005753
                 IPCI
                        A61K0009-16 [ICM, 7]
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                 IPCI
                        A61K [ICM, 7]
     Drugs, especially low aqueous solubility drugs, are provided in a porous
matrix form,
     preferably microparticles, which enhances dissoln. of the drug in aqueous
     media. The drug matrixes preferably are made using a process that
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solubility, in
a volatile solvent to form a drug solution, (ii) combining at least one pore
forming agent with the drug solution to form an emulsion, suspension, or

includes (i) dissolving a drug, preferably a drug having low aqueous

second solns., and (iii) removing the volatile solvent and pore forming agent from the emulsion, suspension, or second solution to yield the porous matrix of drug. The pore forming agent can be either a volatile liquid that is immiscible with the drug solvent or a volatile solid compound, preferably a volatile salt. In a preferred embodiment, spray drying is used to remove the solvents and the pore forming agent. The resulting porous matrix has a faster rate of dissoln. following administration to a patient, as compared to non-porous matrix forms of the drug. In a preferred embodiment, microparticles of the porous drug matrix are reconstituted with an aqueous medium and administered parenterally, or processed using standard techniques into tablets or capsules for oral administration. Paclitaxel or docetaxel can be provided in a porous matrix form, which allows the drug to be formulated without solubilizing agents and administered as a bolus. For example, a nifedipine-loaded organic solution was prepared by dissolving 9.09 g of PEG 3350, 2.27 g of nifedipine, and 0.009 g of lecithin in 182 mL of methylene chloride. An aqueous solution prepared by dissolving 3.27 g of NH4HCO3 and 0.91 g of PEG 3350 in 1.82 mL of water. The aqueous and organic solns. were homogenized and resulting emulsion was spray dried. A suspension of the porous nifedipine drug matrix was prepared in 5% dextrose solution at a concentration of 2.5 mg/mL. A bolus injection of the suspension was tolerated when administrated to dogs. drug solubilization polymer sugar porous matrix; microparticle oral parenteral drug porous matrix Artery Bone Eye Heart Lung Mucous membrane Neoplasm Skin Synovial fluid (administration to; preparation of porous matrixes containing hydrophilic polymers and sugars for enhancement of drug dissoln.) Drug delivery systems (bolus, injections, i.v.; preparation of porous matrixes containing hydrophilic polymers and sugars for enhancement of drug dissoln.) Drug delivery systems (buccal; preparation of porous matrixes containing hydrophilic polymers and sugars for enhancement of drug dissoln.) Drug delivery systems (capsules; preparation of porous matrixes containing hydrophilic polymers sugars for enhancement of drug dissoln.) Estrogens RL: PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (conjugated; preparation of porous matrixes containing hydrophilic polymers sugars for enhancement of drug dissoln.) (conjunctiva, administration to; preparation of porous matrixes containing hydrophilic polymers and sugars for enhancement of drug dissoln.) Drying

was

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polymers

(fluidized-bed; preparation of porous matrixes containing hydrophilic

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and sugars for enhancement of drug dissoln.)
IT
     Pore
        (forming agents; preparation of porous matrixes containing hydrophilic
polymers
        and sugars for enhancement of drug dissoln.)
IT
     Polymers, biological studies
     RL: PEP (Physical, engineering or chemical process); THU (Therapeutic
     use); BIOL (Biological study); PROC (Process); USES (Uses)
        (hydrophilic; preparation of porous matrixes containing hydrophilic
polymers and
        sugars for enhancement of drug dissoln.)
ፐጥ
     Drug delivery systems
       (injections, i.m.; preparation of porous matrixes containing hydrophilic
        polymers and sugars for enhancement of drug dissoln.)
ΙT
     Drug delivery systems
        (injections, i.v.; preparation of porous matrixes containing hydrophilic
        polymers and sugars for enhancement of drug dissoln.)
ΙT
     Drug delivery systems
        (injections, s.c.; preparation of porous matrixes containing hydrophilic
        polymers and sugars for enhancement of drug dissoln.)
IT
     Drug delivery systems
        (intracranial; preparation of porous matrixes containing hydrophilic
polymers
        and sugars for enhancement of drug dissoln.)
TT
     Drug delivery systems
        (intratracheal; preparation of porous matrixes containing hydrophilic
polymers
        and sugars for enhancement of drug dissoln.)
IT
     Drug delivery systems
        (microparticles; preparation of porous matrixes containing hydrophilic
polymers
        and sugars for enhancement of drug dissoln.)
IT
     Drug delivery systems
        (mucosal; preparation of porous matrixes containing hydrophilic polymers and
        sugars for enhancement of drug dissoln.)
IT
     Drug delivery systems
        (nasal; preparation of porous matrixes containing hydrophilic polymers and
        sugars for enhancement of drug dissoln.)
ΙT
     Drug delivery systems
        (oral; preparation of porous matrixes containing hydrophilic polymers and
sugars
        for enhancement of drug dissoln.)
ΙT
     Drug delivery systems
        (parenterals; preparation of porous matrixes containing hydrophilic
polymers and
        sugars for enhancement of drug dissoln.)
TΤ
     Drug delivery systems
        (powders; preparation of porous matrixes containing hydrophilic polymers and
        sugars for enhancement of drug dissoln.)
IT
     Dissolution rate
     Emulsions
     Evaporation
       Freeze drying
     Particle size
     Solubilization
     Surface area
     Suspensions
     Wetting agents
        (preparation of porous matrixes containing hydrophilic polymers and sugars
for
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enhancement of drug dissoln.)
     Interferons
IT
     Interleukins
     Taxanes
     RL: PEP (Physical, engineering or chemical process); THU (Therapeutic
     use); BIOL (Biological study); PROC (Process); USES (Uses)
        (preparation of porous matrixes containing hydrophilic polymers and sugars
for
        enhancement of drug dissoln.)
IT
    Carbohydrates, biological studies
     Lecithins
     Polyoxyalkylenes, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (preparation of porous matrixes containing hydrophilic polymers and sugars
for
        enhancement of drug dissoln.)
TT
    Drug delivery systems
        (rectal; preparation of porous matrixes containing hydrophilic polymers and
        sugars for enhancement of drug dissoln.)
ΙT
     Volatile substances
        (solvents; preparation of porous matrixes containing hydrophilic polymers
and
        sugars for enhancement of drug dissoln.)
TΤ
     Drying
        (spray; preparation of porous matrixes containing hydrophilic polymers and
        sugars for enhancement of drug dissoln.)
TΤ
     Drug delivery systems
        (sublingual; preparation of porous matrixes containing hydrophilic polymers
and
        sugars for enhancement of drug dissoln.)
TT
     Drug delivery systems
        (suppositories, vaginal; preparation of porous matrixes containing
hydrophilic
        polymers and sugars for enhancement of drug dissoln.)
TΤ
     Drug delivery systems
        (suppositories; preparation of porous matrixes containing hydrophilic
polymers
        and sugars for enhancement of drug dissoln.)
TΤ
    Drug delivery systems
        (tablets; preparation of porous matrixes containing hydrophilic polymers and
        sugars for enhancement of drug dissoln.)
ΙT
    Drug delivery systems
        (topical; preparation of porous matrixes containing hydrophilic polymers and
        sugars for enhancement of drug dissoln.)
ΙT
        (vacuum; preparation of porous matrixes containing hydrophilic polymers and
        sugars for enhancement of drug dissoln.)
ΙT
    Drug delivery systems
        (vaginal; preparation of porous matrixes containing hydrophilic polymers and
        sugars for enhancement of drug dissoln.)
TΤ
    Salts, biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (volatile, pore forming agents; preparation of porous matrixes containing
        hydrophilic polymers and sugars for enhancement of drug dissoln.)
TΤ
    Solvents
        (volatile; preparation of porous matrixes containing hydrophilic polymers
and
        sugars for enhancement of drug dissoln.)
TΤ
    631-61-8, Ammonium acetate 1066-33-7, Ammonium bicarbonate 1863-63-4,
    Ammonium benzoate
                         12125-02-9, Ammonium chloride, uses
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RL: NUU (Other use, unclassified); USES (Uses)

(preparation of porous matrixes containing hydrophilic polymers and sugars for enhancement of drug dissoln.) IT 50-28-2, Estradiol, biological studies 50-35-1, Thalidomide Dextrose, biological studies 52-53-9, Verapamil 53-03-2, Prednisone 55-98-1, Busulfan 57-63-6, Ethinyl estradiol 58-61-7, Adenosine, biological studies 59-92-7, Levodopa, biological studies 67-97-0, Vitamin D3 67-97-0D, Vitamin D3, analogs 71 - 58 - 9, Medroxyprogesterone acetate 75-64-9, Erbumine, biological studies 77-36-1, Chlorthalidone 89-57-6, Mesalamine 126-07-8, Griseofulvin 128-13-2, Ursodiol 298-46-4, Carbamazepine 302-79-4, Tretinoin 321-64-2, Tacrine 363-24-6, Dinoprostone 437-38-7, Fentanyl 439-14-5, Diazepam 443-48-1, Metronidazole 518-28-5, Podofilox 745-65-3, Alprostadil 846-49-1, Lorazepam 1951-25-3, Amiodarone 3239-44-9, Dexfenfluramine 4759-48-2, Isotretinoin 5534-09-8, Beclomethasone dipropionate 5593-20-4, Betamethasone dipropionate 9002-68-0, Follitropin 9002-72-6, Growth hormone 9007-12-9, Calcitonin 9041-93-4, Bleomycin sulfate 10238-21-8, Glyburide 11096-26-7, Erythropoietin 12629-01-5, Somatropin 12633-72-6, Amphotericin 13311-84-7, Flutamide 15307-79-6, Diclofenac sodium 15307-86-5, Diclofenac 15687-27-1, Ibuprofen 18559-94-9, Albuterol 20830-75-5, 21256-18-8, Oxaprozin 21829-25-4, Nifedipine Digoxin 22204-53-1, 27203-92-5, Tramadol Naproxen 28860-95-9, Carbidopa 28981-97-7, Alprazolam 29094-61-9, Glipizide 30516-87-1, Zidovudine **32986-56-4**, **Tobramycin** 33069-62-4, Paclitaxel 34911-55-2, Bupropion 36505-84-7, Buspirone 40391-99-9 41340-25-4, Etodolac 41575-94-4, Carboplatin 42399-41-7, Diltiazem 51022-70-9, Albuterol sulfate Nabumetone 51333-22-3, Budesonide 51773-92-3, Mefloquine hydrochloride 54143-55-4, Flecainide 54527-84-3, Nicardipine hydrochloride 54910-89-3, Fluoxetine 54965-21-8, Albendazole 54965-24-1, Tamoxifen citrate 55268-75-2, 56124-62-0, Valrubicin 56180-94-0, Acarbose Cefuroxime 59729-33-8, Citalopram 60142-96-3, Gabapentin 60205-81-4, Ipratropium 63659-18-7, Betaxolol 65277-42-1, Ketoconazole 66085-59-4, Nimodipine 66376-36-1, Alendronate 66852-54-8, Halobetasol propionate 69655-05-6, Didanosine 70476-82-3, Mitoxantrone hydrochloride 72432-03-2, Miglitol 72509-76-3, Felodipine 72558-82-8, Ceftazidime 72956-09-3, Carvedilol 73384-59-5, Ceftriaxone 73590-58-6, Omeprazole 75330-75-5, Lovastatin 75695-93-1, Isradipine 75847-73-3, Enalapril 76095-16-4, Enalapril 76547-98-3, Lisinopril maleate 76824-35-6, Famotidine 76963-41-2. Nizatidine 77883-43-3, Doxazosin mesylate 78246-49-8, Paroxetine hydrochloride 78628-80-5, Terbinafine hydrochloride 78755-81-4. 79517-01-4, Octreotide acetate Flumazenil 79559-97-0, Sertraline 79794-75-5, Loratadine hydrochloride 79902-63-9, Simvastatin 80274-67-5, Metoprolol fumarate 81098-60-4, Cisapride Clarithromycin 82410-32-0, Ganciclovir 82752-99-6, Nefazodone hydrochloride 82834-16-0, Perindopril 83799-24-0, Fexofenadine 83905-01-5, Azithromycin 83919-23-7, Mometasone furoate 84625-61-6, Itraconazole 85721-33-1, Ciprofloxacin 86386-73-4, Fluconazole 86541-74-4, Benazepril hydrochloride 86541-75-5, Benazepril 87679-37-6, Trandolapril 89778-27-8, Toremifene citrate Terbinafine 91421-42-0, Rubitecan 93413-69-5, Venlafaxine 93957-54-1, Fluvastatin 95058-81-4, Gemcitabine 95233-18-4, Atovaquone 97322-87-7, Troglitazone 97048-13-0, Urofollitropin 98048-97-6, Fosinopril 98079-52-8, Lomefloxacin hydrochloride 98319-26-7, Finasteride 99011-02-6, Imiquimod 99294-93-6, Zolpidem tartrate 100286-90-6, Irinotecan hydrochloride 100986-85-4, Levofloxacin 103577-45-3, Lansoprazole 103628-48-4, Sumatriptan succinate

104632-25-9,

103775-10-6, Moexipril 104227-87-4, Famciclovir

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Pramipexole dihydrochloride
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     134678-17-4, Lamivudine
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     Mangafodipir trisodium
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     143011-72-7, Granulocyte colony-stimulating factor
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                   145040-37-5, Candesartan cilexetil
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     154248-97-2, Imiglucerase
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     679809-58-6, Enoxaparin sodium
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        (preparation of porous matrixes containing hydrophilic polymers and sugars
for
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IT
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                                            9003-43-4,
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                            9005-65-6, Tween 80
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            26266-57-9, Span 40
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        (preparation of porous matrixes containing hydrophilic polymers and sugars
for
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IT
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for
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RN
     32986-56-4 HCAPLUS
CN
     D-Streptamine, O-3-amino-3-deoxy-\alpha-D-glucopyranosyl-(1\rightarrow6)-O-
     [2,6-diamino-2,3,6-trideoxy-\alpha-D-ribo-hexopyranosyl-(1\rightarrow4)]-2-
     deoxy- (9CI) (CA INDEX NAME)
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Absolute stereochemistry.

IT 64-17-5, Ethanol, biological studies
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(preparation of porous matrixes containing hydrophilic polymers and sugars for enhancement of drug dissoln.) 64-17-5 HCAPLUS RN Ethanol (9CI) (CA INDEX NAME) CN  $H_3C-CH_2-OH$ L95 ANSWER 7 OF 9 HCAPLUS COPYRIGHT 2006 ACS on STN AN 1999:647601 HCAPLUS DN 132:141845 Entered STN: 12 Oct 1999 ΕD In vitro bactericidal evaluation of a low phase transition temperature ΤI liposomal tobramycin formulation as a dry powder preparation against gram-negative and gram-positive bacteria ΑU Beaulac, C.; Sachetelli, S.; Lagace, J. Department of Microbiology and Immunology Faculty of Medicine, Universite CS de Montreal, Montreal, QC, H3C 3J7, Can. Journal of Liposome Research (1999), 9(3), 301-312 SO CODEN: JLREE7; ISSN: 0898-2104 PB Marcel Dekker, Inc. DT Journal LA English 63-6 (Pharmaceuticals) Section cross-reference(s): 1 In previous studies, delivery of a liquid preparation of encapsulated tobramycin in fluid liposomes, called Fluidosomes, has showed a marked improvement in the bactericidal activity against in-vitro and in-vivo extracellular infections. To examine the possibility of developing aerosol treatment using dehydrated Fluidosomes for the treatment of chronic pulmonary infections, freeze-dried prepns. of tobramycin and Fluidosomes were tested against cultures of Pseudomonas aeruginosa, Stenotrophomonas maltophilia, Burkholderia cepacia, Escherichia coli and Staphylococcus aureus. Bacterial colonies were enumerated 0, 1, 3, 6 and 16 h after the addition of the antibiotic. Sixteen hours post-treatment, the growth of  ${\bf P}.$ aeruginosa, S. maltophilia, B. cepacia and E. coli in the presence of sub-minimal inhibitory concns. of tobramycin was significantly lowered resp. by 17-, 40-, 47-, and 50-fold in comparison with growth in the presence of free antibiotic. No improvement was observed against S. aureus. Results obtained in this study suggest that the dehydrated form of liposomal antibiotic maintains the ability to increase penetration of the antibiotic in gram neg. bacterial cells; the development of aerosolization methods to administer dehydrated liposomes associated with high concns. of antibiotic could be a practical and efficient way of treating chronic pulmonary infections caused by resistant bacteria. STliposome tobramycin bactericide phase transition temp ΙT Antibacterial agents Burkholderia cepacia Escherichia coli Phase transition temperature Pseudomonas aeruginosa Staphylococcus aureus Stenotrophomonas maltophilia (bactericidal evaluation of low-phase transition temperature liposomal tobramycin formulation as dry powder against bacteria)

- krishnan 10 / 827024 IT Phospholipids, biological studies RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (bactericidal evaluation of low-phase transition temperature liposomal tobramycin formulation as dry powder against bacteria) IT 32986-56-4, Tobramycin RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (bactericidal evaluation of low-phase transition temperature liposomal tobramycin formulation as dry powder against bacteria) 2644-64-6, DPPC IT 61361-72-6, Dimyristoylphosphatidylglycerol RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (bactericidal evaluation of low-phase transition temperature liposomal tobramycin formulation as dry powder against bacteria) RE.CNT THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD RF. (1) Ahmed, M; J of Bacteriology 1995, V177, P3904 HCAPLUS (2) Aronoff, S; Pediatric and Pulmonology 1991, V11, P289 MEDLINE (3) Barclay, M; Antimicrobial Agents and Chemotherapy 1992, V36, P1951 HCAPLUS (4) Barclay, M; J of Antimicrobial Chemotherapy 1996, V37, P1155 HCAPLUS (5) Bartlett, G; Clinical and research 1958, V234, P466 (6) Beaulac, C; Antimicrobial Agents and Chemotherapy 1996, V40, P665 HCAPLUS (7) Beaulac, C; J Microencapsulation 1997, V14, P335 HCAPLUS (8) Beaulac, C; J of Antimicrobial Chemother 1998, V41, P35 HCAPLUS (9) Beaulac, C; Master thesis, Universite du Quebec a Trois-Rivieres (UQTR) 1995, P67 (10) Daikos, G; Antimicrobial Agents and Chemotherapy 1991, V35, P117 HCAPLUS (11) Daikos, G; J of Infectious Diseases 1990, V162, P414 HCAPLUS (12) Doit, C; Antimicrobial Agents and Chemotherapy 1994, V38, P2655 HCAPLUS (13) Friedland, I; J of Antimicrobial Chemother 1994, V34, P231 HCAPLUS (14) Gilleland, B; J of Medical Microbiology 1989, V29, P41 (15) Gilleland, H; Canadian Journal of Microbiology 1988, V34, P499 HCAPLUS (16) Grinus, L; J of Biological Chemistry 1994, V269, P29998 (17) Johnson, D; Aerosol Science and Technology 1996, V25, P22 HCAPLUS (18) Lindsay, C; Clinical Pharmacokinetics 1993, V24, P496 MEDLINE (19) Lyon, B; [Review] Microbiological Review 1987, V51, P88 HCAPLUS (20) Mehta, R; Antimicrobial Agents and Chemotherapy 1993, V37, P2584 HCAPLUS (21) National Comittee for Clinical Laboratory Standards; Methods for dilution
- antimicrobial susceptibility tests for bacteria that grow aerobically 1993
- (22) Nikaido, H; Science 1994, V264, P382 HCAPLUS
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- (30) Thomassen, M; American Review of Respiration Diseases 1985, V131, P791 MEDLINE
- (31) Tomasz, A; New England Journal of Medecine 1994, V330, P1247 MEDLINE 32986-56-4, Tobramycin
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(bactericidal evaluation of low-phase transition temperature liposomal tobramycin formulation as dry powder against bacteria)

RN 32986-56-4 HCAPLUS

CN D-Streptamine, O-3-amino-3-deoxy- $\alpha$ -D-glucopyranosyl- $(1\rightarrow6)$ -O-[2,6-diamino-2,3,6-trideoxy- $\alpha$ -D-ribo-hexopyranosyl-(1 $\rightarrow$ 4)]-2-

## deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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L95 ANSWER 8 OF 9 HCAPLUS COPYRIGHT 2006 ACS on STN
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    121:296634
DN
ED
    Entered STN: 24 Dec 1994
    Lyophilized ligand-receptor complexes for assays and sensors
ΤI
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    Ligler, Frances S.; Whelan, James P.
PA
    United States Dept. of the Navy, USA; U.S. Drug Testing, Inc.
SO
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INCL 435005000
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CLASS
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    A dry reagent prepared by lyophilizing a labeled
AB
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     lyophilized support and use of lyophilized beads in a
     flow immunosensor are described, as is a lyophilization ELISA
    plate assay.
ST
     lyophilized reagent ligand receptor complex; sensor
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IT
     Steroids, analysis
     RL: ANT (Analyte); ANST (Analytical study)
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and sensors)
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TT
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ΙT
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ΙT
     Receptors
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (complexes with ligands; lyophilized ligand-receptor
        complexes for assays and sensors)
ΙT
     Blood serum
     Buffer substances and systems
     Surfactants
        (cryoprotectant; preparation of lyophilized ligand-receptor
        complexes for assays and sensors)
IT
     Polysaccharides, biological studies
     Proteins, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (cryoprotectant; preparation of lyophilized liqand-receptor
        complexes for assays and sensors)
TT
     Analysis
     Chromatography, column and liquid
     Immunoassay
     Salmonella
     Sensors
        (lyophilized ligand-receptor complexes for assays and
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IT
     Complement
     Ferritins
     Haptoglobins
     Herbicides
     Histocompatibility antigens
     Insecticides
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     Myoglobins
     Opioids
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     Thyroglobulins
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TΤ
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sensors)
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        (F1, Y. pestis; lyophilized ligand-receptor complexes for
        assays and sensors)
TT
     Immunoglobulins
     RL: ANT (Analyte); ANST (Analytical study)
        (G, lyophilized ligand-receptor complexes for assays and
        sensors)
IT
     Immunoglobulins
     RL: ANT (Analyte); ANST (Analytical study)
        (M, lyophilized ligand-receptor complexes for assays and
        sensors)
TT
     Antigens
     RL: ANT (Analyte); ANST (Analytical study)
        (PSA (prostate-specific antigen), lyophilized ligand-receptor
        complexes for assays and sensors)
TΤ
     Virus, animal
        (adeno-, antigens and antibodies; lyophilized ligand-receptor
        complexes for assays and sensors)
IT
     Toxins
     RL: ANT (Analyte); ANST (Analytical study)
        (anthrax, protein LF (lethal factor), lyophilized
        ligand-receptor complexes for assays and sensors)
IT
     Toxins
     RL: ANT (Analyte); ANST (Analytical study)
        (anthrax, protein PA (protective antigen), lyophilized
```

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ligand-receptor complexes for assays and sensors)
IT
     Receptors
     RL: ANT (Analyte); ANST (Analytical study)
        (cholinergic, antibody; lyophilized ligand-receptor complexes
        for assays and sensors)
IT
     Analysis
        (clin., lyophilized ligand-receptor complexes for assays and
        sensors)
IT
     Ligands
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (complexes, with receptors; lyophilized ligand-receptor
        complexes for assays and sensors)
IT
     Oligosaccharides
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (di-, cryoprotectant; preparation of lyophilized ligand-receptor
        complexes for assays and sensors)
IT
     Toxins
     RL: ANT (Analyte); ANST (Analytical study)
        (endo-, lyophilized ligand-receptor complexes for assays and
        sensors)
IT
     Toxins
     RL: ANT (Analyte); ANST (Analytical study)
        (entero-, lyophilized ligand-receptor complexes for assays
        and sensors)
IΤ
     Metals, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (heavy, lyophilized ligand-receptor complexes for assays and
        sensors)
ΙT
     Virus, animal
        (hepatitis A, antigens and antibodies; lyophilized
        ligand-receptor complexes for assays and sensors)
IT
     Virus, animal
        (hepatitis B, antigens and antibodies; lyophilized
        ligand-receptor complexes for assays and sensors)
TΤ
     Virus, animal
        (herpes, antigens and antibodies; lyophilized ligand-receptor
        complexes for assays and sensors)
IT
     Virus, animal
        (human immunodeficiency, antigens and antibodies; lyophilized
        ligand-receptor complexes for assays and sensors)
TΤ
     Proteins, specific or class
     RL: ANT (Analyte); ANST (Analytical study)
        (p24, lyophilized ligand-receptor complexes for assays and
        sensors)
IT
     Aromatic hydrocarbons, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (polycyclic, lyophilized ligand-receptor complexes for assays
        and sensors)
IT
     Virus, animal
        (rota-, antigens and antibodies; lyophilized ligand-receptor
        complexes for assays and sensors)
TT
     Globulins, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (thyroxine-binding, lyophilized ligand-receptor complexes for
        assays and sensors)
IT
     Fetoproteins
     RL: ANT (Analyte); ANST (Analytical study)
        (α1-, lyophilized ligand-receptor complexes for assays
        and sensors)
```

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ΙT
     Microglobulins
     RL: ANT (Analyte); ANST (Analytical study)
        (β2-, antibody; lyophilized ligand-receptor complexes
        for assays and sensors)
IT
     56-81-5, 1,2,3-Propanetriol, biological studies
                                                      67-68-5, Dimethyl
     sulfoxide, biological studies
                                   25322-68-3
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (cryoprotectant; preparation of lyophilized ligand-receptor
        complexes for assays and sensors)
ΙT
     9002-61-3, Chorionic gonadotropin
     RL: ANT (Analyte); ANST (Analytical study)
        (human; lyophilized ligand-receptor complexes for assays and
        sensors)
IT
     9001-15-4, Creatine kinase
                                 9001-60-9, Lactate dehydrogenase
     RL: ANT (Analyte); ANST (Analytical study)
        (isoenzymes; lyophilized ligand-receptor complexes for assays
        and sensors)
IT
     50-06-6, Phenobarbital, analysis 50-23-7, Cortisol
                                                           50-28-2, Estradiol,
               50-33-9, Phenylbutazone, analysis 50-36-2, Cocaine 50-37-3,
    analysis
    Lysergic acid diethylamide
                                50-49-7, Imipramine
                                                     50-67-9, Serotonin,
    analysis
               50-99-7, Glucose, analysis
                                           51-06-9, Procainamide
    Thyroxine, analysis 52-39-1, Aldosterone
                                                54-36-4, Metyrapone
     55-63-0, Nitroglycerine 56-54-2, Quinidine 56-75-7, Chloramphenicol
     57-27-2, Morphine, analysis 57-41-0, Phenytoin
                                                      57-83-0, Progesterone,
               58-22-0, Testosterone 58-25-3, Chlordiazepoxide 58-55-9,
     analysis
    Theophylline, analysis
                             59-30-3, Folic acid, analysis 60-92-4, Cyclic
          68-19-9, Vitamin B12
                                71-63-6, Digitoxin
                                                    72-44-6, Methaqualone
    76-99-3, Methadone
                                                77-67-8, Ethosuximide
                        77-10-1, Phencyclidine
    78-11-5, Pentaerythritol tetranitrate 88-89-1, Picric acid
    Carbamazine
                  92-52-4D, Biphenyl, chloro derivs.
                                                     99-66-1, Valproic acid
    118-96-7, Trinitrotoluene 121-82-4, Cyclonite 125-33-7, Primidone
    137-58-6, Lidocaine 300-62-9, Amphetamine 439-14-5, Diazepam
    469-62-5, Propoxyphene
                             519-09-5, Benzoylecgonine 525-66-6, Propranolol
    537-46-2, Methamphetamine
                                561-27-3, Heroin 1403-66-3, Gentamicin
    1972-08-3, Tetrahydrocannabinol 6893-02-3, Triiodothyronine 8059-24-3,
                 9001-77-8, Acid phosphatase 9002-62-4, Prolactin, analysis
    Vitamin B6
    9002-64-6, Parathyroid hormone 9002-67-9, Luteinizing hormone
    9002-68-0, FSH
                     9002-71-5, Thyroid-stimulating hormone 9002-72-6,
    Growth hormone
                     9002-76-0, Gastrin
                                         9004-10-8, Insulin, analysis
    9007-92-5, Glucagon, analysis
                                    9015-82-1, Angiotensin converting enzyme
    9035-54-5, Placental lactogen
                                    11096-26-7, Erythropoietin 12794-10-4D,
                                                    30516-87-1, Azidothymidine
    Benzodiazepine, derivs.
                             20830-75-5, Digoxin
    32986-56-4, Tobramycin
                             37221-79-7, Vasoactive
                                                   59112-80-0, C-Peptide
    intestinal polypeptide 37517-28-5, Amikacin
    59763-91-6, Pancreatic polypeptide 59865-13-3, Cyclosporine
    67763-96-6, Somatomedin C 107231-12-9, Botulism toxin
    RL: ANT (Analyte); ANST (Analytical study)
        (lyophilized ligand-receptor complexes for assays and
       sensors)
ΙT
    9014-08-8
    RL: ANT (Analyte); ANST (Analytical study)
        (neuron-specific; lyophilized ligand-receptor complexes for
       assays and sensors)
ΙT
    51-84-3, Acetylcholine, analysis
    RL: ANT (Analyte); ANST (Analytical study)
        (receptor, antibody; lyophilized ligand-receptor complexes
       for assays and sensors)
ΙT
    32986-56-4, Tobramycin
    RL: ANT (Analyte); ANST (Analytical study)
```

(lyophilized ligand-receptor complexes for assays and sensors)

RN 32986-56-4 HCAPLUS

CN D-Streptamine, O-3-amino-3-deoxy- $\alpha$ -D-glucopyranosyl- $(1\rightarrow 6)$ -O-[2,6-diamino-2,3,6-trideoxy- $\alpha$ -D-ribo-hexopyranosyl- $(1\rightarrow 4)$ ]-2-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L95 ANSWER 9 OF 9 HCAPLUS COPYRIGHT 2006 ACS on .STN 1982:612500 HCAPLUS AN 97:212500 DN ED Entered STN: 12 May 1984 TΙ Evaluation of an automated procedure for determining the minimum inhibitory concentrations (MIC): the ABAC MIC ΑU Thabaut, A.; Durosoir, J. L.; Meyran, M. CS Hop. Mil., Saint-Mande, 94160, Fr. SO Pathologie Biologie (1982), 30(6 bis), 555-9 CODEN: PTBIAN; ISSN: 0031-3009 DT Journal LA French 10-5 (Microbial Biochemistry) CC Section cross-reference(s): 9 AB The ABAC system allows for the simultaneous and automatic distribution of a standardized inoculum into microtube cuvettes containing 2-fold serial dilns. of the antibiotic to be tested in lyophilized broth medium. After 18 h incubation, the system automatically prints the MIC. Comparisons were made of the MIC of 7  $\beta$ -lactam antibiotics (ampicillin, carbenicillin, cephalotin, cefoxitin, cefamandole, cefuroxime, and cefotaxime) and 6 aminoglycosides (gentamicin, tobramycin, netilmycin, amikacin, kanamycin, and lividomycin) obtained by the ABAC system and by the agar dilution method for 302 gram-neg. bacteria. A comparison of ABAC and agar dilution methods was made for the MIC of 8 antibiotics (oxacillin, oleandomycin, spiramycin, erythromycin, clindamycin, pristinamycin, doxycycline, and vancomycin) for 117

ST antibiotic min inhibitory concn detn automation

the MICs obtained with the 2 methods is excellent.

IT Antibiotics

(min. inhibitory concns. of, automated determination of)

Staphylococcus aureus strains. The reproductibility of the

=> => fil medline FILE 'MEDLINE' ENTERED AT 11:31:08 ON 06 JUN 2006

results obtained by the ABAC system is good and the correlation between

FILE LAST UPDATED: 3 JUN 2006 (20060603/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 is now (26 Feb.) available. For details on the 2006 reload, enter HELP RLOAD at an arrow prompt (=>). See also:

http://www.nlm.nih.gov/mesh/http://www.nlm.nih.gov/pubs/techbull/nd04/nd04\_mesh.html http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\_med\_data\_changes.html http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\_2006\_MeSH.html

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d all

L131 ANSWER 1 OF 1 MEDLINE on STN

AN 2002212319 MEDLINE

DN PubMed ID: 11948553

- TI Freeze-drying of tert-butanol/water cosolvent systems: a case report on formation of a friable freeze-dried powder of tobramycin sulfate.
- AU Wittaya-Areekul Sakchai; Needham Gregory F; Milton Nathaniel; Roy Michael L; Nail Steven L
- CS Department of Industrial and Physical Pharmacy, School of Pharmacy, 1336 Robert Heine Building, Purdue University, West Lafayette, Indiana 47907, USA.
- SO Journal of pharmaceutical sciences, (2002 Apr) Vol. 91, No. 4, pp. 1147-55.

  Journal code: 2985195R. ISSN: 0022-3549.
- CY United States
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 200208
- ED Entered STN: 12 Apr 2002 Last Updated on STN: 15 Aug 2002 Entered Medline: 14 Aug 2002
- AB A case study is presented in which a tert-butanol (TBA)/water cosolvent system was found to be a useful means of producing freeze-dried tobramycin sulfate that readily forms a loose powder upon agitation in a specialized application in which a critical quality attribute is the ability to pour the sterile powder from the vial. formulation and processing variables are important in achieving acceptable physical properties of the cake as well as minimizing residual TBA levels. Liquid/liquid phase separation was observed above critical concentrations of both drug and TBA, resulting in a two-layered lyophilized cake with unacceptable appearance, physical properties, and residual TBA levels. However, the choice of tobramycin sulfate and TBA concentrations in the single-phase region of the phase diagram resulted in a lyophilized solid that can readily be poured from vials. Crystallization of TBA before drying is critical to achieving adequately low residual TBA levels, and this is reflected in the effect of thermal history of freezing on

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residual TBA levels, where rapid freezing results in incomplete
     crystallization of TBA and relatively high levels of residual solvent.
     Annealing at a temperature above T'(g) of the system after an initial
     freezing step significantly reduces the level of residual TBA. Secondary
     drying, even at increased temperature and for extended times, is not an
     effective method of reducing residual TBA levels.
     Copyright 2002 Wiley-Liss, Inc. and the American Pharmaceutical
     Association J Pharm Sci 91: 1147-1155, 2002
СТ
      Anti-Bacterial Agents: CH, chemistry
      Calorimetry, Differential Scanning
        Freeze Drying: MT, methods
      Powders
      Research Support, Non-U.S. Gov't
      Solutions
       *Solvents: CH, chemistry
        Temperature
       *Tobramycin: CH, chemistry
     *Water: CH, chemistry
       *tert-Butyl Alcohol: CH, chemistry
RN
     32986-56-4 (Tobramycin); 75-65-0 (tert-Butyl Alcohol);
     7732-18-5 (Water)
CN
     0 (Anti-Bacterial Agents); 0 (Powders); 0 (Solutions); 0 (Solvents)
=> => d his
     (FILE 'HOME' ENTERED AT 10:35:25 ON 06 JUN 2006)
                SET COST OFF
     FILE 'REGISTRY' ENTERED AT 10:35:33 ON 06 JUN 2006
                E TOBRAMYCIN/CN
L1
              1 S E3
L2
             24 S C18H37N5O9/MF
L3
             24 S L2 AND OC5/ES AND 46.150.1/RID AND 3/NR
L4
              3 S L3 AND 2 3 6 TRIDEOXY AND 2 DEOXY AND 3 AMINO 3 DEOXY AND STR
L5
              2 S L4 NOT LABELED
             2 S L1, L5
L6
L7
             65 S 66007-88-3/CRN OR 32986-56-4/CRN
\Gamma8
             35 S L7 AND (PMS OR MXS OR IDS)/CI
             30 S L7 NOT L8
L9
L10
             18 S L9 AND COMPD
L11
             12 S L9 NOT L10
L12
             14 S L6, L11
L13
             10 S (METHANOL OR ETHANOL OR 1-PROPANOL OR 2-PROPANOL OR PROPANOL
     FILE 'HCAPLUS' ENTERED AT 10:40:36 ON 06 JUN 2006
L14
           4917 S L12
L15
           4793 S TOBRAMYCIN?
L16
             58 S TOBRAMICIN?
L17
           5447 S L14-L16
L18
             55 S L17 AND L13
L19
              4 S L12(L) PREP+NT/RL AND L18
L20
              2 S L19 NOT (137:30253 OR 102:119707)/DN
                E KWOK/AU
                E KWOK K/AU
L21
             16 S E3, E10, E11
L22
              5 S E30, E31
                E YANG/AU
L23
              4 S E3
                E YANG K/AU
```

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251 S E3
L24
L25
              23 S E23
                 E YANG KANG/AU
             132 S E3,E8
L26
                 E ABRAXIS/PA, CS
                 E AM PHARM/PA, CS
                 E AM PHAR/PA, CS
                 E A PHAR/PA,CS
                 E AME PHAR/PA, CS
                 E AMER PHAR/PA,CS
                 E AMERIC PHAR/PA,CS
                 E AMERICA PHAR/PA, CS
                 E AMERICAN PHAR/PA,CS
L27
               4 S E19-E22
L28
              0 S L17 AND L21-L27
L29
             113 S L17 (L) PREP+NT/RL
             390 S L17 (L) PROC+NT/RL
L30
L31
               9 S L29, L30 AND L18
               5 S L31 NOT L19
L32
L33
               1 S L32 AND 137:358002/DN
L34
               3 S L20, L33
     FILE 'REGISTRY' ENTERED AT 10:49:01 ON 06 JUN 2006
L35
               1 S 75-65-0
     FILE 'HCAPLUS' ENTERED AT 10:49:10 ON 06 JUN 2006
L36
               2 S L35 AND L17
L37
               3 S L34 AND L14-L34, L36
L38
           2193 S L17 AND (P AERUGINOSA? OR PROTEUS? OR P MIRABILIS OR M MORGAN
                 E STAPHYLOCCUS/CT
L39
               7 S E4+OLD, NT
L40
          39556 S E14+OLD, NT
L41
          27095 S E21+OLD, NT
L42
              36 S E23+OLD, NT OR E24+OLD, NT OR E25+OLD, NT
                 E PROVIDENCIA/CT
L43
            938 S E3+OLD, NT
L44
               2 S E15, E16
                 E CITROBACTER/CT
           2742 S E3+OLD, NT
L45
                 E KLEBSIELLA/CT
L46
          10671 S E3+OLD, NT
                 E ENTERBACTER/CT
                 E ENTEROBACTER/CT
           6147 S E3+OLD, NT
L47
                 E SERRATIA/CT
L48
           6134 S E3+OLD, NT
                 E "E COLI"/CT
                 E ESCHERICHIA COLI/CT
         151013 S E3+OLD, NT
L49
                 E PROTEUS/CT
           6724 S E3+OLD, NT OR E5+OLD, NT
L50
L51
               2 S E7
                 E PROTEUS MIRABILIS/CT
L52
           2663 S E3+OLDNT
                 E MORGANELLA MORGANII/CT
            905 S E3+OLD, NT
L53
L54
            395 S E1+OLD, NT
L55
            437 S RETTGERI/CW
                 E PROVIDENCIA RETTGERI/CT
L56
            426 S E3+OLD, NT
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E PSEUDOMONAS RETTGERI/CT
L57
               1 S E3
L58
           22472 S AERUGINOSA/CW
                 E PSEUDOMONAS AERUGINOSA/CT
          21201 S E3+OLD, NT
L59
          16976 S VULGARIS/CW
L60
                 E PROTEUS VULGARIS/CT
           2281 S E3+OLD, NT
L61
L62
            2016 S L17 AND L39-L61
L63
           2439 S L38, L62
                 E SEPTICEMIA/CT
                 E E3+ALL
           7766 S E4+OLD, NT
L64
          28660 S E4/BI OR E6/BI OR E7/BI OR E9/BI OR E10/BI OR E11/BI OR E12/B
L65
                 E UNIARY TRACT INFECTION/CT
                 E URINARY TRACT INFECTION/CT
                 E E3+ALL
L66
            737 S E2
                 E RESPIRATORY INFECTION/CT
                 E E4+ALL
L67
           1557 S E2
                 E SKIN INFECTION/CT
                 E E3+ALL
L68
           1298 S E2,E3
                 E SOFT TISSUE INFECTION/CT
                 E E2+ALL
L69
            168 S E2 (L) INFECT? OR INFECTION?/CT (L) TISSUE(L)SOFT
                 E BURN/CT
                 E E3+ALL
L70
           7966 S E3+OLD, NT
                 E PERITONITIS/CT
L71
            101 S E3+OLD, NT
                 E E3+ALL
L72
            743 S E2,E3
                 E CENTRAL NERVOUS SYSTEM INFECTION/CT
                 E E3+ALL
L73
            336 S E2, E3
L74
            211 S L17 AND L64-L73
            129 S L63 AND L74
L75
           2521 S L63, L74, L75
L76
L77
           2223 S L76 AND L14
L78
               1 S L77 AND LYOPHIL?
L79
              2 S L76 AND LYOPHIL?
L80
              5 S L78, L79, L34
             31 S L76 AND LIQUID
L81
L82
             15 S L76 AND AQUEOUS
L83
             10 S L76 AND (FREEZ? OR FROZ? OR VACUUM?)
L84
               5 S L81, L82 AND L83
                 SEL DN 1 3 5
L85
               2 S L84 NOT E1-E3
               6 S L80, L85
L86
L87
               6 S L86 AND L14-L34, L36-L86
                 E FREEZE DRYING/CT
L88
           6932 S E3+OLD, NT
                 E SOLVENT/CT
          56978 S E64+OLD, NT
L89
L90
             11 S L17 AND L88
L91
               4 S L90 AND L89
L92
               3 S L90 AND L18
L93
               5 S L91, L92
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L94
               4 S L93 NOT 136:391077/DN
1.95
               9 S L87, L94 AND L14-L34, L36-L94
                 SEL HIT RN
     FILE 'REGISTRY' ENTERED AT 11:11:02 ON 06 JUN 2006
L96
               4 S E1-E4
     FILE 'REGISTRY' ENTERED AT 11:11:23 ON 06 JUN 2006
     FILE 'HCAPLUS' ENTERED AT 11:11:31 ON 06 JUN 2006
     FILE 'WPIX' ENTERED AT 11:13:44 ON 06 JUN 2006
            480 S L15 OR L16
1.97
                E TOBRAMYCIN/CN
               3 S E3-E7
L98
            447 S (RA7VYL OR R03158 OR R02067)/DCN OR 2067/DRN OR (170288-1-0-0
1,99
            610 S L97, L99
L100
L101
              8 S (METHANOL OR ETHANOL OR 1-PROPANOL OR 2-PROPANOL OR PROPANOL
                 SEL SDCN
                EDIT /SDCN /DCN
          15286 S E1-E8
L102
L103
          45015 S (0245 OR 0270 OR 0271 OR 0302 OR 0304 OR 0373 OR 0431 OR 0436
                 SEL L101 DCSE
                EDIT /DCSE /DCRE
          10170 S E9-E16
L104
              9 S L100 AND L102-L104
L105
L106
             18 S L100 AND (B10-E04 OR C10-E04 OR B10-E04D OR C10-E04D)/MC
T-107
              0 S L100 AND E10-E04/MC
L108
              0 S L100 AND (E10-E04E? OR E10-E04F)/MC
L109
              2 S L100 AND A61K009-19/IPC, IC, ICM, ICS, ICA, ICI
L110
             54 S L100 AND A61K009-14/IPC, IC, ICM, ICS, ICA, ICI
L111
             10 S L100 AND (B12-M11G OR C12-M11G)/MC
L112
              5 S L105, L106 AND L109-L111
L113
             14 S L106, L106 NOT L112
T.114
             54 S L110, L111 NOT L112-L113
     FILE 'MEDLINE' ENTERED AT 11:27:10 ON 06 JUN 2006
           4900 S L12 OR L15 OR L16
L115
L116
              2 S L115 AND L13
                E SOLVENT/CT
L117
         327529 S E4+NT
                E TERT-BUTYL ALCOHOL/CT
L118
            201 S E3+NT
L119
             28 S L115 AND L117, L118
L120
             28 S L116, L119
                E FREEZE DRYING/CT
L121
              1 S E3+NT AND L120
                E TEMPERATURE/CT
1.122
              2 S E3+NT AND L120
L123
              2 S L121, L122
L124
           2371 S L115 AND B3./CT
                E BACTERIAL INFECTION/CT
           1719 S E6+NT AND L115
L125
L126
           3081 S L124, L125
L127
             11 S L120 AND L126
L128
              1 S L123 AND L127
L129
              1 S L123 NOT L128
L130
              2 S ANTI-BACTERIAL AGENTS+NT/CT AND L123
L131
              1 S L129 AND L130
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FILE 'MEDLINE' ENTERED AT 11:31:08 ON 06 JUN 2006